

ADME NTP Study S0533 Formamide

The contract laboratory abbreviation for the test article is FMD.

Species: young adult male F344 rats and B6C3F1 mice.

Vehicles: intravenous, sterile 0.9% saline; inhalation, conditioned room air.

CASRN 75-12-7

Radiolabeled with carbon-14; [¹⁴C]Formamide

Formamide Studies Performed:

- Single intravenous 10 mg [¹⁴C]FMD/kg dose in rats with sacrifice 24 hours postdose. (Study A; Tables 1, 2)
- Nose-only inhalation exposure of 20 ppm [¹⁴C]FMD for 6 hours in rats with sacrifice immediately after exposure and at 66 hour following termination of exposure. (Study B; Tables 3,4)
- Single intravenous 10 mg [¹⁴C]FMD/kg dose in mice with sacrifice 72 hours postdose. (Study C; Table 5,6)
- Single intravenous 10 mg [¹⁴C]FMD/kg dose in rats and mice with sacrifice 72 hours postdose. (Study E, Clearance of radioactivity from blood; Tables 7,8,9,10)
- Single intravenous 10 mg [¹⁴C]FMD/kg dose in mice 4 hours following a pretreatment with 100 mg/kg 1-aminobenzotriazole (ABT) that was administered by intraperitoneal injection. Mice were sacrificed 24 hours postdose. (Study F; Table 11)
- Single intravenous 10 mg [¹⁴C]FMD/kg dose in mice following inhibition of Cytochrome P450 isozyme 2E1, all P450, or no inhibition with sacrifice 72 hours postdose. (Study G; Table 12,13,14)
- Single intravenous 10 mg [¹⁴C]FMD/kg dose in mice following inhibition of Cytochrome P450 enzyme 2E1 with sacrifice 24 hours postdose. (Study H; Table 13)

The bile excretion study (Study D) had no tables. In Study E, rats were jugular vein cannulated for serial blood samples and four mice were sampled per time point with the blood from two mice pooled (yielding only two samples per timepoint). The general Cytochrome P450 inhibition (with ABT) and specific P450 isozyme 2E1 inhibition with trans-1,2-dichloroethylene (DCE) of Study G was achieved with a single intraperitoneal administration of ABT (100 mg/kg) or DCE (472 mg/kg) 4 hours or 2 hours prior to formamide administration, respectively. To achieve inhibition throughout Study H, 575

mg/kg DCE was administered every 4 hours with the first administration 4 hours prior to the 10 mg [¹⁴C]FMD/kg intravenous dose.

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Table 1

Execution of Total ¹⁴C over 24 h by Male Rats Administered a Single iv Dose of [¹⁴C]FMD
(All Values Expressed as % of Administered Dose)^a

End of Collection (h)	Volatile Breath	CO ₂ Breath	Urine	Cage Rinse	Feces	Carcass & Tissues	Total Recovered Dose
3	0 ^b	3.9 ± 0.4	c	c	c	c	3.9 ± 0.4
6	0.1 ± 0.0 ^b	5.3 ± 0.6	15.5 ± 1.7	c	c	c	20.9 ± 2.2
12	0.1 ± 0.0	9.0 ± 0.7	9.2 ± 1.6	c	0.4 ± 0.2	c	18.7 ± 1.8
24	0.19 ± 0.2	10.7 ± 0.9	14.6 ± 2.1	1.4 ± 0.3	0.7 ± 0.2	19.1 ± 2.9	46.8 ± 3.9
Overall Mean Recovery	0.4 ± 0.3	26.9 ± 1.7	38.9 ± 1.5	1.4 ± 0.3	1.1 ± 0.3	19.1 ± 2.9	90.2 ± 0.7

^a Mean ± S.D. (N=5)

^b Value less than 0.05.

^c No samples collected.

Table 2

Concentration and Tissue to Blood Ratios of Total ¹⁴C in Excised Tissues of Male Rats 24 h
Post Administration of a Single iv Dose of [¹⁴C]FMD

Tissue	Concentration (µg eq/g)			Tissue to Blood Ratio		
		±			±	
Adipose	0.32	±	0.04	0.12	±	0.02
Bladder	3.850	±	1.08	1.39	±	0.37
Blood	2.77	±	0.20		Unity	
Brain	2.06	±	0.25	0.74	±	0.05
Carcass	0.63	±	0.07	0.23	±	0.01
Cecal Contents	3.25	±	0.57	1.17	±	0.15
Cecum	0.11	±	0.02	0.04	±	0.01
Heart	2.47	±	0.27	0.891	±	0.04
Small Intestine	0.119	±	0.03	0.04	±	0.01
Kidney	2.74	±	0.28	0.99	±	0.05
Large Intestine	0.24	±	0.08	0.08	±	0.03
Large Intestine Contents	3.52	±	0.59	1.27	±	0.15
Liver	2.43	±	0.28	0.88	±	0.07
Lung	2.48	±	0.27	0.90	±	0.04
Muscle	2.02	±	0.32	0.73	±	0.09
Small Intestine Contents	2.70	±	0.58	0.97	±	0.16
Spleen	2.68	±	0.28	0.97	±	0.04
Skin	1.84	±	0.22	0.66	±	0.06
Testes	2.23	±	0.42	0.80	±	0.11

^aN=5

Table 3

Radioactivity Excreted Over Time from Male Rats Receiving a Single Inhaled [¹⁴C]FMD Dose

(All Values Expressed as % of Administered Dose)^a

End of Collection (h)	Volatile Breath	CO ₂ Breath	Urine	Cage Rinse	Feces	Carcass & Tissues	Total Recovered Dose
6	0.1 ± 0.0 ^b	7.6 ± 0.6	2.5 ± 0.5	d	d	d	10.1 ± 1.0
18	0.2 ± 0.1	9.8 ± 0.9	5.1 ± 1.9	d	0.5 ± 0.2	d	15.6 ± 2.7
42	0.3 ± 0.3	6.4 ± 0.3	8.8 ± 1.9	d	0.9 ± 0.2	d	16.4 ± 2.0
66	0.1 ± 0.1	3.3 ± 0.2	7.8 ± 1.1	6.1 ± 1.1	0.5 ± 0.1	39.4 ± 4.6	57.3 ± 5.0
Overall Mean Recovery	0.7 ± 0.5	27.1 ± 1.3	24.6 ± 3.6	6.1 ± 1.1	2.1 ± 0.3	39.4 ± 4.6	100 ^c

^a All values expressed as Mean ± S.D. (N=5)

^b Value less than 0.05.

^c Administered dose is calculated as the absorbed dose (total radioactivity in the carcass and tissues and all excreta).

^d No sample collected.

Table 4

Concentration of Radiolabel and Tissue to Blood Ratios of Total ^{14}C in Excised Tissues^b from Male Rats Sacrificed Immediately (top) and 66 h (bottom) Post Administration of a Single Inhaled Dose of [^{14}C]FMD^a

Tissue	Concentration ($\mu\text{g eq/g}$)			Tissue to Blood Ratio		
Adipose	0.15	\pm	0.04	0.11	\pm	0.03
Bladder	1.95	\pm	0.40	1.46	\pm	0.32
Blood	1.35	\pm	0.17	Unity		
Brain	1.04	\pm	0.17	0.77	\pm	0.06
Carcass	1.55	\pm	0.25	1.15	\pm	0.11
Cecum	0.09	\pm	0.01	0.06	\pm	0.00
Heart	1.18	\pm	0.15	0.88	\pm	0.03
Small Intestine	0.12	\pm	0.02	0.09	\pm	0.01
Kidney	1.55	\pm	0.24	1.15	\pm	0.06
Large Intestine	0.03	\pm	0.00	0.02	\pm	0.00
Liver	1.28	\pm	0.18	0.95	\pm	0.07
Lung	1.32	\pm	0.19	0.98	\pm	0.11
Muscle	1.11	\pm	0.30	0.82	\pm	0.16
Plasma	1.52	\pm	0.28	1.13	\pm	0.13
Spleen	1.27	\pm	0.16	0.95	\pm	0.02
Stomach	0.05	\pm	0.01	0.04	\pm	0.01
Testes	1.03	\pm	0.12	0.77	\pm	0.04

Tissue	Concentration ($\mu\text{g eq/g}$)			Tissue to Blood Ratio		
Adipose	0.06	\pm	0.02	0.09	\pm	0.01
Bladder	0.54	\pm	0.08	0.86	\pm	0.13
Blood	0.64	\pm	0.18	Unity		
Brain	0.29	\pm	0.08	0.46	\pm	0.08
Carcass	0.74	\pm	0.17	1.17	\pm	0.10
Cecum	0.03	\pm	0.01	0.05	\pm	0.01
Heart	0.46	\pm	0.12	0.72	\pm	0.07
Small Intestine	0.04	\pm	0.01	0.06	\pm	0.01
Kidney	0.51	\pm	0.13	0.81	\pm	0.06
Large Intestine	0.02	\pm	0.00	0.04	\pm	0.01
Liver	0.49	\pm	0.13	0.77	\pm	0.08
Lung	0.51	\pm	0.11	0.80	\pm	0.08
Muscle	0.36	\pm	0.16	0.56	\pm	0.17
Plasma	0.38	\pm	0.11	0.60	\pm	0.09
Spleen	0.55	\pm	0.15	0.86	\pm	0.03
Stomach	0.02	\pm	0.01	0.02	\pm	0.00
Testes	0.30	\pm	0.09	0.48	\pm	0.10

^a N=5

^b The ears were collected as representative skin samples. However, these samples appeared to be highly contaminated due to direct deposition of radiolabel from the FMD atmosphere onto the ears. Radioactivity associated with the ear samples was not included in the calculation of absorbed dose.

Table 5

Radioactivity Excreted Over Time from Male Mice Administered a Single iv [¹⁴C]FMD Dose(All Values Expressed as % of Administered Dose)^a

End of Collection (h)	Volatile Breath	CO ₂ Breath	Urine	Cage Rinse	Feces	Carcass & Tissues	Total Recovered Dose
3	0.3 ± 0.1	20.8 ± 3.3	c	c	c	c	21.1 ± 3.2
6	0.3 ± 0.1	15.0 ± 1.0	10.0 ± 11.5	c	c	c	25.2 ± 10.9
12	0.3 ± 0.1	12.9 ± 1.8	5.6 ± 11.0	c	0.9 ± 0.1	c	19.6 ± 12.0
24	0.3 ± 0.1 _b	3.8 ± 0.6	11.9 ± 10.4	c	1.1 ± 0.4	c	17.0 ± 11.0
48	0.2 ± 0.0 _b	0.6 ± 0.1 _b	1.6 ± 0.5	c	0.4 ± 0.4	c	2.73 ± 0.8
72	0.1 ± 0.0 _b	0.2 ± 0.0 _b	0.7 ± 0.1	0.9 ± 0.3	0.1 ± 0.1	2.8 ± 0.1	4.7 ± 0.3
Overall Mean Recovery	1.4 ± 0.1	53.2 ± 4.1	29.7 ± 1.3	0.9 ± 0.3	2.5 ± 0.3	2.8 ± 0.1	89.7 ± 5.4

^a All values expressed as Mean ± S.D. (N=4)^b Value less than 0.05.^c No samples collected.

Table 6

Concentration and Tissue to Blood Ratios of Total ¹⁴C in Excised Tissues of Male Mice 72 h Post Administration of a Single IV Dose of [¹⁴C]FMD^a

Tissue	Concentration			Tissue to Blood Ratio		
	(µg eq/g)					
Adipose	0.07	±	0.01	0.06	±	0.01
Bladder	0.29	±	0.01	0.23	±	0.02
Blood	1.26	±	0.10	Unity		
Brain	0.15	±	0.01	0.12	±	0.00 ^b
Carcass	0.03	±	0.00 ^b	0.02	±	0.00 ^b
Cecum	0.01	±	0.00 ^b	0.01	±	0.00 ^b
Heart	0.52	±	0.07	0.42	±	0.05
Small Intestine	0.01	±	0.00 ^b	0.01	±	0.0 ^b
Kidney	0.43	±	0.03	0.34	±	0.01
Large Intestine	0.01	±	0.00 ^b	0.00 ^b	±	0.00 ^b
Liver	0.34	±	0.04	0.27	±	0.02
Lung	0.56	±	0.08	0.44	±	0.06
Muscle	0.23	±	0.02	0.18	±	0.01
Plasma	0.14	±	0.02	0.11	±	0.01
Skin	0.21	±	0.01	0.17	±	0.01
Spleen	0.49	±	0.04	0.40	±	0.06
Stomach	0.00 ^b	±	0.00 ^b	0.00 ^b	±	0.00 ^b
Testes	0.09	±	0.01 ^b	0.08	±	0.00 ^b

^aN=4

^b Value less than 0.005.

Table 7

**Concentration of ¹⁴C in Plasma after Administration of
10 mg FMD/kg iv to Male F-344 Rats
(μ g-Equivalents per gram Plasma)**

a. Group A

Sample ^a Time (h)	ER-A1	ER-A2	ER-A3	ER-A4	Mean \pm SD
0	0	0	0	0	0
0.5	12.2	14.1	12.8	13.4	13.1 \pm 0.8
1.5	11.8	12.8	12.1	12.5	12.3 \pm 0.4
3	11.1	11.6	11.3	11.2	11.3 \pm 0.2
8	8.50	8.87	8.69	8.56	8.65 \pm 0.16
24	3.13	3.20	3.11	2.72	3.04 \pm 0.22

b. Group B

Sample ^a Time (h)	ER-B1	ER-B2	ER-B3	ER-B4	Mean \pm SD
0.25	14.8	14.3	14.2	10.1	13.4 \pm 2.2
1	12.8	12.6	11.7	10.1	11.8 \pm 1.2
2	11.7	11.7	11.4	9.90	11.2 \pm 0.9
5	10.2	10.2	10.3	8.42	9.80 \pm 0.92
12	7.55	7.43	7.37	5.95	7.07 \pm 0.75
32	2.02	2.29	1.84	1.74	1.97 \pm 0.24
48	0.670	0.867	0.679	0.670	0.722 \pm 0.097
72	0.204	0.259	0.237	0.208	0.227 \pm 0.026

Average ¹⁴C Concentration (μ g-eq/g) in Plasma

Sample Time (h) ^a	AVG μ g-eq/g Plasma
0	0
0.25	13.4
0.5	13.1
1	11.8
1.5	12.3
2	11.2
3	11.3
5	9.80
8	8.65
12	7.07
24	3.04
32	1.97
48	0.722
72	0.227

^a Times following administration of radiolabeled dose.

Table 8

**Concentration of ¹⁴C in Blood after Administration of
10 mg FMD/kg iv to Male F-344 Rats^a
(μ g-Equivalents per gram Blood)**

a. Group A

Sample ^b Time (h)	ER-A1	ER-A2	ER-A3	ER-A4	Mean \pm SD
0	0	0	0	0	0
0.5	11.0	12.8	11.6	12.1	11.9 \pm 0.7
1.5	10.7	11.5	10.8	11.1	11.0 \pm 0.4
3	9.96	10.4	10.2	10.1	10.2 \pm 0.2
8	7.92	8.11	7.93	7.86	7.96 \pm 0.11
24	3.12	3.07	3.03	2.67	2.97 \pm 0.20

b. Group B

Sample ^a Time (h)	ER-B1	ER-B2	ER-B3	ER-B4	Mean \pm SD
0.25	12.9	13.0	12.8	9.05	11.9 \pm 1.9
1	11.8	11.1	10.9	9.22	10.7 \pm 1.1
2	10.4	10.4	10.2	8.83	9.94 \pm 0.74
5	9.04	8.97	9.14	7.76	8.73 \pm 0.65
12	6.94	6.64	6.76	5.60	6.48 \pm 0.60
32	2.20	2.45	2.17	2.00	2.21 \pm 0.19
48	1.03	1.31	1.10	1.03	1.12 \pm 0.13
72	0.712	0.766	0.804	0.693	0.744 \pm 0.051

Average ¹⁴C Concentration (μ g-eq/g) in Blood Summary

Sample Time (h) ^a	AVG μ g-eq/g Blood
0	0
0.25	11.9
0.5	11.9
1	10.7
1.5	11.0
2	9.94
3	10.2
5	8.73
8	7.96
12	6.48
24	2.97
32	2.21
48	1.12
72	0.744

^a Using the average hematocrit reading found in 24 & 72 h blood samples, the concentration in blood was calculated by adding 66.8% of the plasma concentration/timepoint plus 33% of the RBC concentration/timepoint. Example: ER-A1 0.5h, Plasma Conc. = 12.2 μ g-eq/g, RBC Conc. = 8.70 μ g-eq/g, Blood Conc. = (12.2 x 0.668) + (8.70 x 0.333) = 11.0 μ g-eq/g Blood.

^b Times following administration of radiolabeled dose.

Table 9

Concentration and Plasma to Blood Ratio of Total ¹⁴C in Male Rats and Mice
after iv Administration of 10 mg FMD/kg^a

a. F-344 Rats

Timepoint (h) ^b	Tissue Name	N	ng-eq /g Tissue	Plasma:Blood Ratio
24	Blood	4	2970 ± 203	1.02 ± 0.02
	Plasma	4	3040 ± 218	
32	Blood	4	2205 ± 186	0.89 ± 0.04
	Plasma	4	1973 ± 241	
48	Blood	4	1119 ± 132	0.64 ± 0.02
	Plasma	4	722 ± 97	
72	Blood	4	744 ± 51	0.31 ± 0.02
	Plasma	4	227 ± 26	

b. B6C3F₁ Mice

Timepoint (h) ^b	Tissue Name	N	ng-eq /g Tissue	Plasma:Blood Ratio
24	Blood	2 ^c	687	0.336
	Plasma		231	
48	Blood	2 ^d	968 ^e	0.163
	Plasma		159 ^e	
72	Blood	2 ^c	1170	0.115
	Plasma		135	

^a Study E data.

^b Times following administration of radiolabeled dose.

^c Each of the two determinations were made on the combined blood from two animals (total number of animals = 4)

^d One of the two determinations were made on the combined blood from the two animals that received the rat dosing solution. Doses for these animals were 42.5 and 49.2 mg FMD/kg. The other determination was from a single animal that received a dose of 7.30 mg FMD/kg.

^e Due to problems in dosing, two mice in the 48 h group of Study E received doses of FMD 4-5 times as large as the planned 10 mg FMD/kg dose (Refer to Table 10.1); the ng-eq/g tissue values shown were normalized to a dose of 10 mg/kg.

Table 10

Urinary Excretion of ¹⁴C by Male B6C3F₁ Mice following
a Single 10 mg/kg iv Dose of FMD (Study E)^a

End of Collection	Percent Dose Excreted ^b			Cumulative Percent Dose Excreted ^b		
6 h	17.5	±	4.6	17.5	±	4.5
12 h	3.9	±	3.3	21.4	±	2.6
24 h	6.7	±	2.2	28.1	±	2.1
48 h	2.3	±	1.1	30.4	±	1.8
72 h	0.9	±	0.3	31.3	±	1.8

^a N=4

^b All values expressed as Mean ± S.D.

Table 11

Elimination of $^{14}\text{CO}_2$ by Male B6C3F₁ Mice Following a Single 10 mg/kg iv Dose of FMD
(Study F)^a Administered 4h After a Single 100 mg/kg ABT ip Dose

End of Collection	Percent Dose Excreted ^b			Cumulative Percent Dose Excreted ^b		
3 h	1.4	±	0.1	1.4	±	0.1
6 h	0.5	±	0.1	1.9	±	0.2
12 h	1.2	±	0.2	3.1	±	0.3
24 h	2.2	±	0.9	5.3	±	0.9

^a N=4

^b All values expressed as Mean ± S.D.

Table 12
Elimination of ¹⁴C in Urine by Male B6C3F₁ Mice Administered a Single 10 mg/kg FMD iv Dose^a

(a) Control Animals^b

End of Collection	Percent Dose Excreted		Cumulative Percent Dose Excreted		
6 h	0.0 ±	0.0	0.0 ±	0.0	
12 h	14.38 ±	7.58	14.38 ±	7.58	
24 h	5.34 ±	2.35	19.72 ±	9.86	
48 h	5.77 ±	2.95	25.49 ±	7.12	
72 h	1.83 ±	2.18	27.33 ±	5.40	

(b) FMD Administration 5 h after a Single 100 mg/kg ABT ip Dose^c

End of Collection	Percent Dose Excreted		Cumulative Percent Dose Excreted		
6 h	12.08 ±	12.08	12.08 ±	12.08	
12 h	11.95 ±	7.02	24.03 ±	17.14	
24 h	23.47 ±	10.97	47.50 ±	6.16	
48 h	10.04 ±	3.31	57.54 ±	2.86	
72 h	4.36 ±	1.27	61.90 ±	1.59	

(c) FMD Administration 3 h after a Single 472 mg/kg DCE ip Dose^c

End of Collection	Percent Dose Excreted		Cumulative Percent Dose Excreted		
6 h	2.42 ±	1.83	2.42 ±	1.83	
12 h	6.46 ±	1.65	8.87 ±	0.18	
24 h	5.55 ±	3.29	14.42 ±	3.12	
48 h	4.87 ±	2.11	19.28 ±	5.24	
72 h	1.40 ±	2.62	20.68 ±	7.86	

(d) FMD Administration 3 h after a Single 545 mg/kg DCE ip Dose with 5 Additional DCE Doses Every 4 h^d

End of Collection	Percent Dose Excreted		Cumulative Percent Dose Excreted		
6 h	4.08 ±	8.03	4.08 ±	8.033	
12 h	0.84 ±	1.65	4.92 ±	9.68	
24h	0.84 ±	0.95	5.76 ±	10.04	

^a All Values are expressed as percent of ¹⁴C dose.

^b Values are means ± SD, N=3

^c Values are averages ± range, N=2. Absorption of urine by a food occlusion in the metabolism chamber of one animal from this group was suspected prior to observing significantly low urinary ¹⁴C excretion for that animal. The animal's data is not represented here.

^d Values are means ± SD, N=4

Table 13

Elimination of $^{14}\text{CO}_2$ by Male B6C3F₁ Mice Administered a Single 10 mg/kg FMD iv Dose^a
Studies G and H

(a) control animals^b

End of Collection	Percent Dose Excreted		Cumulative Percent Dose Excreted	
3 h	19.16 ±	2.69	19.16 ±	2.69
6 h	5.70 ±	6.16	24.88 ±	8.86
12 h	11.29 ±	2.35	36.17 ±	11.08
24 h	13.15 ±	8.43	49.32 ±	3.24

(b) FMD Administration 5h After a Single 100 mg/kg ABT ip Dose^b

End of Collection	Percent Dose Excreted		Cumulative Percent Dose Excreted	
3 h	2.27 ±	0.33	2.27 ±	0.33
6 h	0.48 ±	0.10	2.75 ±	0.43
12 h	0.92 ±	0.07	3.67 ±	0.47
24 h	1.47 ±	0.66	5.14 ±	1.11

(c) FMD Administration 3h After a Single 472 mg/kg DCE ip Dose^b

End of Collection	Percent Dose Excreted		Cumulative Percent Dose Excreted	
3 h	3.09 ±	0.56	3.09 ±	0.56
6 h	6.93 ±	2.33	10.02 ±	2.89
12 h	21.55 ±	3.26	31.58 ±	1.99
24 h	17.13 ±	4.39	48.70 ±	2.67

(d) FMD Administration 3h After a Single 545 mg/kg DCE ip Dose with 5 Additional DCE Doses every 4h^c

End of Collection	Percent Dose Excreted		Cumulative Percent Dose Excreted	
3h	1.67 ±	0.34	1.67 ±	0.34
6 h	0.93 ±	0.07	2.60 ±	0.32
12 h	1.57 ±	0.22	4.17 ±	0.50
24 h	2.15 ±	0.94	6.31 ±	1.26

^a All values expressed as Mean ± S.D.

^b N=3

^c N=4.

Table 14

Concentration and Tissue to Blood Ratios of Total ¹⁴C in Excised Tissues of Male Mice 72 h Post Administration of a Single IV Dose of [¹⁴C]FMD

Control Mice

Tissue	Concentration (µg eq/g)			Tissue to Blood Ratio		
Plasma	0.10	±	0.00	0.14	±	0.04
Blood	0.80	±	0.26	Unity		
Liver	0.29	±	0.06	0.37	±	0.04
Muscle	0.21	±	0.02	0.27	±	0.05

Mice Pretreated with ABT ca. 5 h Prior to FMD Administration

Tissue	Concentration (µg eq/g)			Tissue to Blood Ratio		
Plasma	0.11	±	0.07	0.48	±	0.27
Blood	0.23	±	0.05	Unity		
Liver	0.23	±	0.05	1.01	±	0.05
Muscle	0.15	±	0.03	0.658	±	0.121

Mice Pretreated with a Single Dose of DCE ca. 3 h Prior to FMD Administration

Tissue	Concentration (µg eq/g)			Tissue to Blood Ratio		
Plasma	0.16	±	0.02	0.15	±	0.03
Blood	1.13	±	0.12	Unity		
Liver	0.54	±	0.06	0.49	±	0.09
Muscle	0.37	±	0.00	0.33	±	0.04

^a All values expressed as Mean ± S.D., N=3.